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Previously we identified MIR16 (membrane interacting protein of RGS16) as an integral membrane glycoprotein that interacts with ***regulator***

signaling proteins and shares

protein

of

G

significant sequence homology with bacterial ***glycerophosphodiester**

phosphodiesterases

Es), suggesting that it is a putive

mammalian GDE. Here we show that MIR16 belongs to a large, evolutionarily ***glycerophosphodiester*** conserved family of GDEs with a characteristic putative catalytic domain that shares a common motif (amino acids 92-116) with the catalytic domains of mammalian phosphoinositide phospholipases C. Expression of wild-type MIR16 (renamed GDE1), but not two catalytic domain mutants (E97AD99A and H112A), leads to a dramatic increase in glycerophosphoinositol phosphodiesterase (GPI-PDE) activity in HEK 293T cells. Analysis of substrate specificity shows that GDE1MIR16 selectively hydrolyzes GPI over glycerophosphocholine. The GPI-PDE activity of GDE1MIR16 expressed in HEK 293T cells can be regulated by stimulation of G protein-coupled, alpha beta-adrenergic, and lysophospholipid receptors. Membrane topology studies suggest a model in which the catalytic GDE domain faces the lumenextracellular space and the C terminus faces the cytoplasm. Our results suggest that by serving as a PDE for GPI with its activity ***G*** regulated by ***protein*** ***signaling*** , GDE1MIR16 provides a link between phosphoinositide metabolism and G protein signal transduction.

L6 ANSWER 2 OF 3 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2000:240374 BIOSIS DOCUMENT NUMBER: PREV200000240374

TITLE: MIR16, a putative membrane glycerophosphodiester

phosphodiesterase, interacts with RGS16.

AUTHOR(S): Zheng, Bin; Chen, Dan; Farquhar, Marilyn Gist (1)

CORPORATE SOURCE: (1) Department of Cellular and Molecular Medicine,

University of California San Diego, La Jolla, CA,

92093-0651 USA

SOURCE: Proceedings of the National Academy of Sciences of the

United States of America, (April 11, 2000) Vol. 97, No. 8,

pp. 3999-4004.

ISSN: 0027-8424.

DOCUMENT TYPE: Article
LANGUAGE: English
SUMMARY LANGUAGE: English

We have identified the protein MIR16 (for Membrane Interacting protein of RGS16) from a yeast two-hybrid screen by using RGS16 as bait. MIR16 shares strong homology with bacterial glycerophosphodiester phosphodiesterases. It interacts with RGS16 and, more weakly, with several other selected RGS proteins. Analysis of deletion mutants showed that the N-terminal region of the RGS domain in RGS16 is required for its interaction with MIR16. MIR16 is an integral membrane glycoprotein, because it remained associated with membrane fractions after alkaline treatment and because, in some cells, it is sensitive to digestion with endoglycosidase H. By immunofluorescence and immunoelectron microscopy, MIR16 was localized on the plasma membrane in liver and kidney and on intracellular membranes in rat pituitary and cultured pituitary cells. MIR16 represents the only integral membrane protein identified thus far to interact with an RGS domain and, to our knowledge, is the only mammalian glycerophosphodiester phosphodiesterase that has been cloned. The putative enzymatic activity of MIR16 and its interaction with RGS16 suggest that it may play important roles in lipid metabolism and in G protein signaling.

L6 ANSWER 3 OF 3 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2002:155792 BIOSIS DOCUMENT NUMBER: PREV200200155792

TITLE: MIR16, a membrane glycerophosphodiester phosphodiesterase:

Enzymatic activity, membrane topology and implications for

membrane trafficking and signaling.

AUTHOR(S): Zheng, Bin (1); Peters, Eugenia; Williams, Chester;

Ferraris, Joan; Burg, Maurice; Schmieder, Sandra (1)
CORPORATE SOURCE: (1) Department of Cellular and Molecular Medicine,

University of California San Diego, 9500 Gilmar Dr.,

CMM-West, Rm 218, La Jolla, CA, 92093-0651 USA

Molecular Biology of the Cell, (Dec., 2000) Vol. 11, No.

Supplement, pp. 485a. http://www.molbiolcell.org/. print. Meeting Info.: 40th American Society for Cell Biology Annual Meeting San Francisco, CA, USA December 09-13, 2000

ISSN: 1059-1524.

DOCUMENT TYPE: Conference LANGUAGE: English

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52 S GLYCEROPHOSPHORYL DIESTER PHOSPHODIESTERASE

170 S L1 OR L2

1882 S REGULATOR (P) (G PROTEIN SIGNALING)

6 S L1 (P) L4

3 DUPLICATE REMOVE L5 (3 DUPLICATES REMOVED)

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L1

L2 L3

L4

L5

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